

**AMENDMENTS****To the specification:**

On page 14, please replace the paragraph starting on line 27 with the following:

Figure 8A and 8B is a combination graph showing the purification of ts149 by anion exchange chromatography on PI matrix, eluted with a linear 900-1300 meq NaCl gradient at pH 8.0.

On page 15, please replace the paragraph starting on line 3 with the following:

Figure 10A and 10B is a combination graph showing the separation of Adenovirus and recombinant AAV. The upper panel shows separation on PI anion-exchange matrix, eluted with a 0-1000 meq gradient of NaCl at pH 8.0. The lower panel shows subsequent separation of Adenovirus from contaminants on HS cation-exchange matrix, eluted with a 0-500 meq gradient of NaCl at pH 8.0.

On page 15, please replace the paragraph starting on line 8 with the following:

Figure 11A and 11B is two bar graphs, showing the effect of fetal bovine serum levels (FBS) in the culture medium on rAAV production. The upper graph indicates DRPs; the lower graph indicates RUs. Serum deficiency in the culture medium is one of a number of stress factors that the producer cells can be subjected to in order to enhance the production of viral particles.

On page 17, please replace the paragraph starting on line 9 with the following:

Figure 26 shows the relationship of 26 A and 26 B.

Figure 26A and 26 B provides amino acid and vitamin compositions for the media supplements described in Example 17.

On page 106, please replace the paragraph starting on line 11 with the following:

The results are shown in Figures 31A, 31B [-] and 32. As shown in Figure 31A, the majority of DRPs (70%) were released into the media with adjustment of the osmolality and

corresponding conductivity at day 2 of rAAV production and that adjustment of the osmolality to 450 mOsm and conductivity with NaCl 4 hours prior to harvest on day 3 results in a majority of DRPs released in the cell culture medium (56%). A significant increase in release of rAAV vector was observed for all conditions in which osmolality (and thus conductivity) was adjusted after day 0 as compared to control.

On page 109, please replace the paragraph starting on line 11 with the following:

As indicated in Figures 33A, 33B and 34A and 34 B, the increase in rAAV vector released into culture medium for NaCl formulated cultures as opposed to sorbitol formulated cultures was not due to a significant difference in metabolic rate (glucose consumption), total cell density, or osmolality of the cultures as there were no appreciable differences in the cultures. However, the cultures did differ in their conductivity (Table 12). Compare, for example, the NaCl formulation at a starting osmolality of 250 mOsm with a conductivity of 10.00 mS that increased to 13.29 mS by day 3 to the sorbitol 300 mOsm starting formulation with a conductivity of 7.11 mS that increased to 10.29 mS by day 3. Generally, the NaCl formulations demonstrating a range of conductivities between approximately 10 and 15 mS demonstrate the greatest percentage release of rAAV vector into the supernatant.